Fig. 1a

Stage I

Undifferentiated embryonic stem cells cultured on feeder cells

Stage II (7 days)

Generation of nesting positive islet cell progenitors in suspended embryoid bodies

Stage III (7 days)

Culture of embryoid bodies in ITSF medium: elimination of nestin negative cells/progenitor isolation

Stage IV (7 days)

Culture of progenitors with N2, B27, bFGF supplementation: expansion of progenitors

V (14 days)

Replacement of bFGF with nicotinamide and reduction of glucose concentration to 5 mM: generation of surface-bound islet like clusters containing insulin producing cells

Stage VI (14 days)

Suspension culture of dissociated surface-bound clusters:

- generation of purified suspended islet like clusters
- steep increase in insulin secretion capacity
- purification of insulin producing cells
- prolongation of in-vitro longevity of insulin producing cells





Fig. 1c

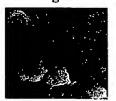


Fig. 1d



Fig. 1e



Fig. 1f



Fig. 1g



Fig. 2

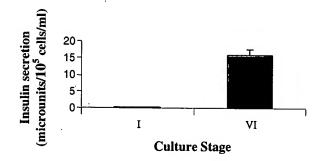


Fig. 3

STAGES

ES III IV V-H V-L VI

Insulin	
Somatostatin	
PDX1	(1773) = r
ngn3	
Glucagon	
Pax4	
Pax6	Consult.
Isl1	
Nkx6.1	fa .a.s.
Glut2	House s
GAPDH	

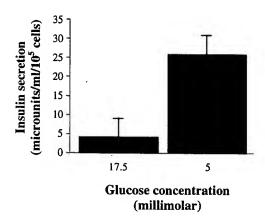


Fig. 4

Fig. 5a

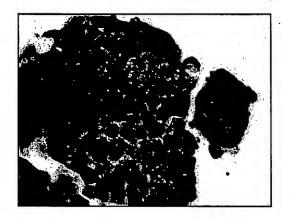


Fig. 5b

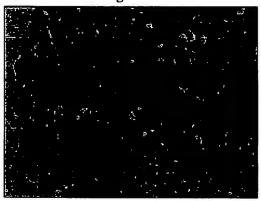
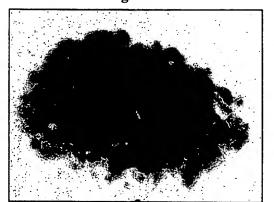


Fig. 5c



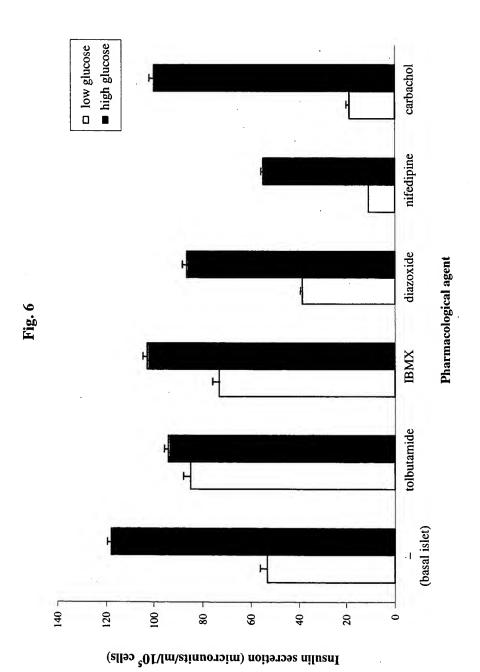
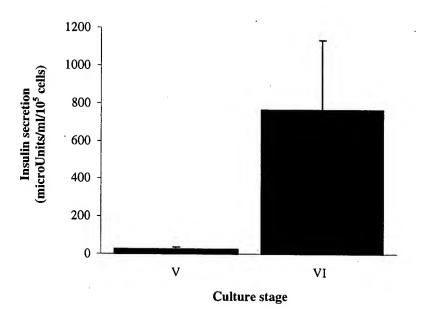
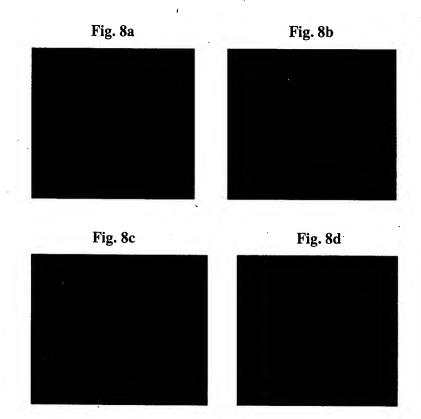


Fig. 7



8/10



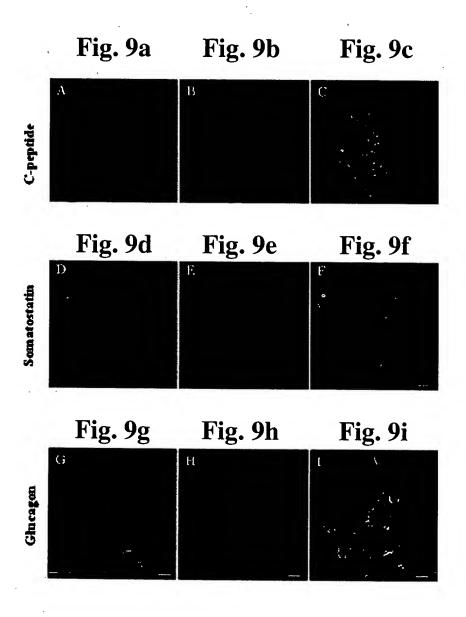


Fig. 10

